, TENT COOPERATION TREATY

NOTIFICATION OF THE RECORDING OF A CHANGE (PCT Rule 92bis.1 and Administrative Instructions, Section 422) Date of mailing (day/month/year) 11 August 2000 (11.08.00) Applicant's or agent's file reference PHM 70478/WO International application No. PCT/GB00/00357 International filing date (day/month/year) 08 February 2000 (08.02.00) 1. The following indications appeared on record concerning:		From the INTERNATIONAL BUREAU
AstraZenaca Global Intellectual Property Mereside, Alderley Park Macclesfield Cheshire SK10 4GR ROYAUME-UNI Date of mailing (day/month/year) 11 August 2000 (11.08.00) Applicant's or agent's file reference PHM 70478/WO International application No. PCT/GB00/00357 International filing date (day/month/year) 08 February 2000 (08.02.00) 1. The following indications appeared on record concerning:	PCT	То:
ROYAUME-UNI	OF A CHANGE (PCT Rule 92bis.1 and	AstraZenaca Global Intellectual Property Mereside, Alderley Park Macclesfield
Applicant's or agent's file reference PHM 70478/WO International application No. PCT/GB00/00357 International filing date (day/month/year) 08 February 2000 (08.02.00) 1. The following indications appeared on record concerning: X the applicant the inventor the agent the common representative		
International application No. PCT/GB00/00357 International filing date (day/month/year) 08 February 2000 (08.02.00) 1. The following indications appeared on record concerning: X		
PCT/GB00/00357 08 February 2000 (08.02.00) 1. The following indications appeared on record concerning: X		IMPORTANT NOTIFICATION
Name and Address ASTRAZENECA UK LIMITED 15 Stanhope Gate London W1Y 6LN United Kingdom 2. The International Bureau hereby notifies the applicant that the following change has been recorded concerning: the person X the name X the address X the nationality X the residence ASTRAZENECA AB S-151 85 Södertälje Sweden Teleprinter No. State of Nationality X the residence SE SE Telephone No. Teleprinter No. Teleprinter No. Teleprinter No. Teleprinter No. Teleprinter No.		
Name and Address ASTRAZENECA UK LIMITED 15 Stanhope Gate London W1Y 6LN United Kingdom 2. The International Bureau hereby notifies the applicant that the following change has been recorded concerning: the person X the name X the address X the nationality X the residence Name and Address ASTRAZENECA AB S-151 85 Södertälje Sweden ASTRAZENECA AB S-151 85 Södertälje Sweden		, the again.
United Kingdom Facsimile No. Teleprinter No. 2. The International Bureau hereby notifies the applicant that the following change has been recorded concerning: the person X the name X the address X the nationality X the residence Name and Address ASTRAZENECA AB S-151 85 Södertälje Sweden State of Nationality State of Residence SE SE Telephone No. Teleprinter No.	ASTRAZENECA UK LIMITED	GB GB
2. The International Bureau hereby notifies the applicant that the following change has been recorded concerning: the person X the name X the address X the nationality X the residence	London W1Y 6LN United Kingdom	Facsimile No.
Name and Address ASTRAZENECA AB S-151 85 Södertälje Sweden X the name X the address X the nationality X the residence SE SE Telephone No. Facsimile No. Teleprinter No.		Teleprinter No.
Name and Address ASTRAZENECA AB S-151 85 Södertälje Sweden Facsimile No. Teleprinter No.		
S-151 85 Södertälje Sweden Facsimile No. Telephone No. Teleprinter No.		State St. Laws.
Teleprinter No.	S-151 85 Södertälje	Telephone No.
		Facsimile No.
3. Further observations, if necessary:		Teleprinter No.
	3. Further observations, if necessary:	
4. A copy of this notification has been sent to:	4. A copy of this notification has been sent to:	
X the receiving Office X the designated Offices concerned		X the designated Offices concerned
the International Searching Authority the elected Offices concerned		the elected Offices concerned
the International Preliminary Examining Authority other:		other:
	The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland	Authorized officer Mougamadou ABIDINE
The International Bureau of WIPO 34, chemin des Colombettes Mougamadou ABIDINE	1	

LATENT COOPERATION TREATY

	From the INTERNATIONAL BUREAU		
PCT	То:		
NOTIFICATION OF THE RECORDING OF A CHANGE (PCT Rule 92bis.1 and Administrative Instructions, Section 422) Date of mailing (day/month/year) 11 August 2000 (11.08.00)	BILL, Kevin AstraZenaca Global Intellectual Property Mereside, Alderley Park Macclesfield Cheshire SK10 4GR ROYAUME-UNI		
Applicant's or agent's file reference	IMPORTANT NOTIFICATION		
PHM 70478/WO			
International application No. PCT/GB00/00357	International filing date (day/month/year) 08 February 2000 (08.02.00)		
The following indications appeared on record concerning: the applicant the inventor	the agent the common representative		
Name and Address	State of Nationality State of Residence		
BILL, Kevin Global Intellectual Property AstraZeneca UK Limited Mereside, Alderley Park	Telephone No. +44 1625 512461		
Macclesfield Cheshire SK10 4TG United Kingdom	Facsimile No. +44 1625 583358		
Office Kingdom	Teleprinter No.		
2. The International Bureau hereby notifies the applicant that the	e following change has been recorded concerning:		
the person the name X the add			
Name and Address	State of Nationality State of Residence		
BILL, Kevin AstraZenaca Global Intellectual Property Mereside, Alderley Park	Telephone No. +44 1625 512461		
Macclesfield Cheshire SK10 4GR	Facsimile No. +44 1625 583358		
United Kingdom	Teleprinter No.		
3. Further observations, if necessary:			
4. A copy of this notification has been sent to:			
X the receiving Office	X the designated Offices concerned		
the International Searching Authority	the elected Offices concerned		
the International Preliminary Examining Authority	other:		
The International Bureau of WIPO	Authorized officer		
34, chemin des Colombettes 1211 Geneva 20, Switzerland	Mougamadou ABIDINE		
Facsimile No.: (41-22) 740.14.35	Telephone No.: (41-22) 338.83.38		

HATENT COOPERATION TREATY

	From the INTERNATIONAL BUREAU			
PCT	То:			
NOTIFICATION OF ELECTION	Assistant Commissioner for Patents United States Patent and Trademark Office			
(PCT Rule 61.2)	Box PCT Washington, D.C.20231 ETATS-UNIS D'AMERIQUE			
Date of mailing (day/month/year) 04 October 2000 (04.10.00)	in its capacity as elected Office			
International application No. PCT/GB00/00357	Applicant's or agent's file reference PHM 70478/WO			
International filing date (day/month/year) 08 February 2000 (08.02.00)	Priority date (day/month/year) 11 February 1999 (11.02.99)			
Applicant				
LITTLE, Stephen et al				
The designated Office is hereby notified of its election made: X in the demand filed with the International Preliminary Examining Authority on:				
30 August 2000 (30.08.00)				
in a notice effecting later election filed with the International Bureau on:				
2. The election X was was not made before the expiration of 19 months from the priority Rule 32.2(b).	date or, where Rule 32 applies, within the time limit under			

Authorized officer The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland Facsimile No.: (41-22) 740.14.35

Zakaria EL KHODARY

Telephone No.: (41-22) 338.83.38

RECEIVED

1 2 MAY 2000

ASTRA JENECA FLO

PATENT COOPERATION TREATY

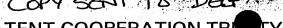


INTELLECTUAL PROPERTY I	From t	ne INTERNATIONAL E	BUREAU
PCT	To:		
NOTIFICATION OF THE RECORDING OF A CHANGE (PCT Rule 92bis.1 and Administrative Instructions, Section 422) Date of mailing (day/month/year) 05 May 2000 (05.05.00)	Glob Astro Merc Mac Ches	, Kevin pal Intellectual Propert aZeneca UK Limited eside, Alderley Park clesfield shire SK10 4TG AUME-UNI	ty
Applicant's or agent's file reference			
PHM 70478/WO		IMPORTANT NOT	TIFICATION
International application No. PCT/GB00/00357		nal filing date (day/month/) ebruary 2000 (08.02.0	•
The following indications appeared on record concerning: The applicant the inventor	the agen	t the comm	on representative
Name and Address		State of Nationality GB	State of Residence GB
ZENECA LIMITED 15 Stanhope Gate London W1Y 6LN United Kingdom		Telephone No.	J GB
		Facsimile No.	
	:	Teleprinter No.	
2. The International Bureau hereby notifies the applicant that t	he following	change has been recorded	concerning:
X the person the name the add	dress	the nationality	the residence
Name and Address		State of Nationality GB	State of Residence
ASTRAZENECA UK LIMITED 15 Stanhope Gate London W1Y 6LN United Kingdom		Telephone No.	GB
Officea Kingdoff		Facsimile No.	
	•	Teleprinter No.	
3. Further observations, if necessary:			
4. A copy of this notification has been sent to:			
X the receiving Office	ŗ	the designated Offices	concerned
X the International Searching Authority	Ī	the elected Offices con	cerned
the International Preliminary Examining Authority		other:	
The International Bureau of WIPO	Authorized of		11
34, chemin des Colombettes 1211 Geneva 20, Switzerland		Lazar Joseph	Papakal

Facsimile No.: (41-22) 740.14.35

Telephone No.: (41-22) 338.83.38

3.



To:

NT COOPERATION TR

2 1 AUG 2000

E RECORDING

From the INTERNATIONAL BUREAU

BILL, Kevin AstraZenaca **Global Intellectual Property** Mereside, Alderley Park Macclesfield Cheshire SK10 4GR

(PCT Rule 92bis.1 and Administrative Instructions, Section 422)

OF A CHANGE

Date of mailing (day/month/year) 11 August 2000 (11.08.00)	ROY	AUME-UNI	
Applicant's or agent's file reference PHM 70478/WO		IMPORTANT NOT	IFICATION
International application No.	Internatio	nal filing date (day/month/y	ear)
PCT/GB00/00357	08 F	ebruary 2000 (08.02.00	D)
1. The following indications appeared on record concerning: X the applicant the inventor Name and Address ASTRAZENECA UK LIMITED 15 Stanhope Gate London W1Y 6LN	the ager	state of Nationality GB Telephone No.	on representative State of Residence GB
United Kingdom		Facsimile No.	
		i acsimile ivo.	
		Teleprinter No.	
242		बर्द्धाः	
2. The International Bureau hereby notifies the applicant that the the person X the name X the add		the nationality	concerning: X the residence
Name and Address		State of Nationality	State of Residence
ASTRAZENECA AB S-151 85 Södertälje		SE Telephone No.	SE
Sweden		relephone No.	
		Facsimile No.	
		Teleprinter No.	
3. Further observations, if necessary:			
4. A copy of this notification has been sent to:	- : 		
X the receiving Office	ſ	X the designated Offices	concerned
the International Searching Authority	Ī	the elected Offices con	cerned
the International Preliminary Examining Authority		other:	
	Authorized	officer	
The International Bureau of WIPO 34, chemin des Colombettes		= 00K	ou ABIDINE —

Telephone No.: (41-22) 338.83.38

Form PCT/IB/306 (March 1994)

Facsimile No.: (41-22) 740.14.35

1211 Geneva 20, Switzerland

003462402



PCT

INTERNATIONAL PRELIMINARY EXAMINATION REPORT

(PCT Article 36 and Rule 70)

			· 			
Applicant's or agent's file reference See Notification of Transmittal of International Preliminary Examination Report (Form PCT/IPEA/416)						
PHM 70478/WO						
	application No.	International filing date (day/month	ı/year)	•	day/month/yea `	ar)
	0/00357 94	08/02/2000		11/02/1999	,	
International C12Q1/68	Patent Classification (IPC) or na	ational classification and IPC				
Applicant						
ASTRAZE	ENECA AB et al					
		ination report has been prepared	t by this Inter	rnational Prel	iminary Exar	mining Authority
1. This in and is	ternational preliminary exam transmitted to the applicant a	according to Article 36.	a by this inter	mational i rei	iriiriai y Exai	
		-				
2. This R	EPORT consists of a total of	8 sheets, including this cover s	heet.			
□ Th	nis report is also accompanie	ed by ANNEXES, i.e. sheets of the sis for this report and/or sheets of	ne description	n, claims and/	'or drawings ade before th	which have
be (s	een amended and are the bar ee Rule 70.16 and Section 6	07 of the Administrative Instructi	ons under the	PC UDE	DATE	
				CODE	DATE	NTD
These annexes consist of a total of sheets.						
				REC'D 2	6 MAR 20	01 GIPS
O Thinks	annet contains indications rela	ating to the following items:		DATA		
3. This report contains indications relating to the following iter				ENTERED	DEL	r.L.
Ⅰ				FINAL		7
II □ Priority				CHECK	***************************************	
III Non-establishment of opinion with regard to novel			ventive step a	and industrial	applicability	
IV.	Lack of unity of inventi	on	. 11		industrial an	nliochilit <i>u</i>
V	Reasoned statement u citations and explanati	under Article 35(2) with regard to ions suporting such statement	noveity, inve	ntive step or	industriai ap	phcability,
l vi	☐ Certain documents cit					
VII		international application				
VIII Certain observations on the international application						
Date of subi	mission of the demand	Date of	completion of	this report		
Euro or out				,		
30/08/200	00	22.03.2	2001			
		A. Albari	and officer			
Name and r	nailing address of the internation examining authority:	ai Authori	zed officer			NO ACCUES PARCULANT
promining,	European Patent Office					(Name of the state
<i>()</i>	D-80298 Munich Tel. +49 89 2399 - 0 Tx: 52365	Filkori 56 epmu d	n, A-C			
_			one No. +49 89	2399 8688		11304C - 30.

INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No. PCT/GB00/00357

I. Basis of the report

1.	resp the	oonse to an invitation	on under Article 14 are referred to in this report as "originally filed" and are not annexed to lo not contain amendments (Rules 70.16 and 70.17).):
	1-45	5	as originally filed
	Clai	ms, No.:	
	1-16	3	as originally filed
	Dra	wings, sheets:	
	1/4-	4/4	as originally filed
	Seq	uence listing part	t of the description, pages:
	1-2,	as originally filed	
2.	With	n regard to the lang Juage in which the	guage, all the elements marked above were available or furnished to this Authority in the international application was filed, unless otherwise indicated under this item.
	The	se elements were	available or furnished to this Authority in the following language: , which is:
		the language of a	translation furnished for the purposes of the international search (under Rule 23.1(b)).
		the language of p	ublication of the international application (under Rule 48.3(b)).
		the language of a 55.2 and/or 55.3).	translation furnished for the purposes of international preliminary examination (under Rule
3.	With	n regard to any nuo rnational prelimina	cleotide and/or amino acid sequence disclosed in the international application, the ry examination was carried out on the basis of the sequence listing:
	\boxtimes	contained in the ir	nternational application in written form.
	Ø	filed together with	the international application in computer readable form.
		furnished subsequ	uently to this Authority in written form.
		furnished subsequ	uently to this Authority in computer readable form.
		The statement that the international a	at the subsequently furnished written sequence listing does not go beyond the disclosure in application as filed has been furnished.
		The statement the	at the information recorded in computer readable form is identical to the written sequence urnished.

4. The amendments have resulted in the cancellation of:

INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No. PCT/GB00/00357

		the description,	pages:
		the claims,	Nos.:
		the drawings,	sheets:
5.		This report has beer considered to go be	n established as if (some of) the amendments had not been made, since they have been yound the disclosure as filed (Rule 70.2(c)):
		(Any replacement si report.)	heet containing such amendments must be referred to under item 1 and annexed to this
6.	Add	litional observations,	if necessary:
			opinion with regard to novelty, inventive step and industrial applicability
1.	The obv	questions whether the questions, or to be industi	ne claimed invention appears to be novel, to involve an inventive step (to be non- rially applicable have not been examined in respect of:
		the entire internation	nal application.
	\boxtimes	claims Nos. 5.	
be	caus	se:	
		the said internationa not require an interr	al application, or the said claims Nos. relate to the following subject matter which does national preliminary examination (<i>specify</i>):
	⊠	the description, clai that no meaningful o see separate shee	ms or drawings (<i>indicate particular elements below</i>) or said claims Nos. 5 are so unclear opinion could be formed (<i>specify</i>):
		the claims, or said could be formed.	claims Nos. are so inadequately supported by the description that no meaningful opinion
		no international sea	arch report has been established for the said claims Nos
2.	and	neaningful internation d/or amino acid seque tructions:	nal preliminary examination report cannot be carried out due to the failure of the nucleotide ence listing to comply with the standard provided for in Annex C of the Administrative
		the written form has	s not been furnished or does not comply with the standard.
			able form has not been furnished or does not comply with the standard.
۷.	Re	asoned statement u	under Article 35(2) with regard to novelty, inventive step or industrial applicability;

citations and explanations supporting such statement

INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No. PCT/GB00/00357

1. Statement

Novelty (N) Yes: Claims 3,6-14

No: Claims 1,2,4,15,16

Inventive step (IS) Yes: Claims 6-14

No: Claims 1-4,15,16

Industrial applicability (IA) Yes: Claims 1-4,6-16

No: Claims -

2. Citations and explanations see separate sheet

VII. Certain defects in the international application

The following defects in the form or contents of the international application have been noted: see separate sheet

VIII. Certain observations on the international application

The following observations on the clarity of the claims, description, and drawings or on the question whether the claims are fully supported by the description, are made: see separate sheet

EXAMINATION REPORT - SEPARATE SHEET

Re Item III

Since there is no sequence information as to the genomic DNA and its origin in claim 5, it is impossible for the skilled person to determine the scope of the claim. Hence, claim 5 contravenes Art 5 PCT and no opinion can be formed on novelty, inventiveness and industrial applicability of said claim (Art 34(4)(a)(ii) PCT).

Re Item V

The following documents are referred to in this communication:

D1: WO 97 31256 A D2: EP-A-0 799 897 D3: WO 93 25563 A

The method according to the present application involves targeting polynucleotides composed of two parts: one part can hybridize to the target molecule and the second part can bind to a capture oligonucleotide which is immobilized on a solid support. The methods rely firstly on the binding of the target molecule comprised in a sample to the targeting polynucleotide and secondly on capturing the hybrid molecule by hybridization with the immobilized capture oligonucleotide.

The capture oligonucleotides are arranged in arrays consisting of a plurality of immobilized capture oligonucleotides of different sequences and melting temperatures within a range of 16°C.

- Novelty (Art 33(2) PCT): 1
- 1.1 Claim 1 is not novel in view of each of documents D2 (p 4 l 54-56: capture oligonucleotides at pre-defined positionally distinct sites and p 4 I 35-45) and D3 (p 7 para 3; Example IV: p 14-16). The same applies to claims 2 and 4 (D2: p 5 l 1-7; D3: Fig 3).
- 1.2 Claim 15 is anticipated by each of D1 (D1: claim 138, 145) and D2 (D2: p 5 | 8-14; p 4 I 35-45; claims 23 and 24). D2 also destroys the novelty of claim 16. Instructions for use are not considered to be a technical feature.

EXAMINATION REPORT - SEPARATE SHEET

- 1.3 Claim 6 is novel, because none of the available documents describes an assay design embracing targeting polynucleotides and capture oligonucleotides that are immobilized on a solid support.
 - For the same reason also claims 7-14 are novel.
- Inventive Step (Art 33(3) PCT): 2
- Claim 3 does not appear to satisfy Art 33(3) PCT. Labelling the immobilized 2.1 capture probes does not appear to solve a problem, because it is not apparent how the discrimination between captured and non captured target sequences can be accomplished (see appl. p 22 I 7-10), when only all the capture oligonucleotides are labelled.
- 2.2 Claim 6 appears to comply with Art 33(3) PCT for the following reasons: D1, which is considered to represent the closest prior art, describes a method which involves two targeting oligonucleotides and immobilized capture oligonucleotides. The subject-matter of claim 6 is distinguished from the method of D1 in that according to D1 a ligation step connects a labelled target specific oligonucleotide with an adjacent target-specific oligonucleotide that additionally contains a portion which is complementary to immobilized capture oligonucleotides, whereas the method according to claim 6 obviates the ligation step and provides for the detection by localization of the hybridized targeting oligonucleotide on the solid support.

The problem to be solved by the method of claim 6 can thus be regarded as the provision of an alternative method for detecting the presence or absence of test nucleic acid sequences in a sample.

None of the available documents renders the subject-matter of claim 6 obvious. Claims 7-9 seem to be inventive, accordingly.

- Claim 10 seems to satisfy Art 33(3) PCT for the following reasons:
 - D1 is considered to represent the closest prior art.
 - The subject-matter of claim 10 is distinguished from the disclosure of D1 in that according to claim 10 the detection is accomplished by incorporating a label into the cDNA during reverse transcription. Hence, again, the ligation step as required

for the method of D1 is omitted.

The problem solved by claim 10 can thus be regarded as the provision of an alternative detection method.

None of the available documents seems to render the subject-matter of claim 10 obvious. The same argument applies to the dependent claims 11-13 and independent claim 14.

Re Item VII

Contrary to the requirements of Rule 5.1(a)(ii) PCT, the relevant background art disclosed in the documents D1, D2 and D3 is not mentioned in the description, nor are these documents identified therein.

Re Item VIII

- Claim 1 does not satisfy Art 6 PCT, because the expression "substantially the 1 same melting temperature" is misleading and renders the scope of the claim unclear. According to the description, "substantially the same melting temperature" is supposed to mean the melting temperature of the average Tm of all immobilized oligonucleotides ±8°C (p 5 I 10-14). This temperature range in the context of specific hybridization experiments does clearly not allow the designation "substantially the same temperature". In order to overcome this objection, the cited explanation given in the description (p 5 I 10-14) should have been incorporated into the claim.
 - Similarly, independent claims 6, 10, 14 and 15 need amendment, since the term "substantially" leads to uncertainty as to the scope for which protection is sought.
- Claim 5 does not satisfy Art 6 PCT, because the scope of the claim is not clearly 2 defined, since it is not clear, as to what "genomic DNA" refers, because no origin of the genomic DNA is given and for most organisms the genomic DNA sequences are not yet known in their entirety.
- Claim 14 lacks an essential feature and hence does not satisfy Art 6 PCT: 3 According to the application, the capture oligonucleotides immobilized on the solid support should each possess the same melting temperature (Tm) when

INTERNATIONAL PRELIMINARY

International application No. PCT/GB00/00357

EXAMINATION REPORT - SEPARATE SHEET

hybridized to their complementary sequences in particular where quantitative analyses is required like in the method of claim 14 (see p 25 I 26 - p 26 I 4). As each independent claim should contain all the essential technical features of the invention, said feature should have been introduced into claim 14.

	For receiving Office use only
International Applic	eation No.
International Filing	Date
	Office and "PCT International Application"

REQUEST	International Filing Date			
The undersigned requests that the present international application be processed according to the Patent Cooperation Treaty.	Name of receiving Office and "PCT International Application"			
-	Applicant's or agent's file (if desired) (12 characters i	e reference maximum) PHM 70478/WO		
Box No. I TITLE OF INVENTION				
NOVEL MATRIX				
Box No. II APPLICANT				
Name and address: (Family name followed by given name; for a legal e The address must include postal code and name of country. The country of Box is the applicant's State (that is, country) of residence if no State of re	entity, full official designation. f the address indicated in this esidence is indicated below.)	This person is also inventor.		
ZENECA Limited		Telephone No.		
15 Stanhope Gate London		+44-1625-516173		
WIY 6LN		Facsimile No.		
		+44-1625-583358		
		Teleprinter No. 669095/669388		
State (that is, country) of nationality:	State (that is, country			
GB		GB		
This person is applicant for the purposes of: all designated the United States all designated the United States		e United States the States indicated in the Supplemental Box		
Box No. III FURTHER APPLICANT(S) AND/OR (FUR	THER) INVENTOR(S)			
Name and address: (Family name followed by given name; for a legal of The address must include postal code and name of country. The country of Box is the applicant's State (that is, country) of residence if no State of re	entity, full official designation. of the address indicated in this esidence is indicated below.)	This person is:		
LITTLE, Stephen		applicant only		
Gadbrook Park Northwich		applicant and inventor		
Cheshire GB				
		inventor only (If this check-box is marked, do not fill in below.)		
State (that is, country) of nationality:	State (that is, country) of residence: GB		
GB				
for the purposes of: States the United	States of America of	e United States America only the States indicated in the Supplemental Box		
Further applicants and/or (further) inventors are indicated on a continuation sheet.				
Box No. IV AGENT OR COMMON REPRESENTATIV	E; OR ADDRESS FOR	CORRESPONDENCE		
The person identified below is hereby/has been appointed to act of the applicant(s) before the competent International Authorities	s as:	gent common representative		
Name and address: (Family name followed by given name; for a legal The address must include postal code and name	entity, full official designation. of country.)	Telephone No. +44-1625-512461		
BILL, Kevin				
Global Intellectual Property AstraZeneca UK Limited		Facsimile No.		
Mereside, Alderley Park, Macclesfield		+44-1625-583358		
Cheshire. SK10 4TG - GB		Teleprinter No.		
	···	669095/669388		
Adress for correspondence: Mark this check-box where no agent or common representative is/has been appointed and the space above is used instead to indicate a special address to which correspondence should be sent.				

Sheet No. 2

Continuation of Box No. III FURTHER APPLICANTS AND/OR (FURTHER) INVENTORS				
If none of the following sub-boxes is used, this sheet should not be included in the request.				
Name and address: (Family name followed by given name; for a legal en The address must include postal code and name of country. The country of a Box is the applicant's State (that is, country) of residence if no State of residence GIBSON, Neil James Gadbrook Park Northwich Cheshire GB	ity, full official designation. the address indicated in this dence is indicated below.) This person is: applicant only applicant and inventor inventor only (If this check-box is marked, do not fill in below.)			
State (that is, country) of nationality: GB	State (that is, country) of residence: GB			
This person is applicant all designated all designated	States except the United States the States indicated in			
for the purposes of: States the United Sta				
Name and address: (Family name followed by given name; for a legal ent The address must include postal code and name of country. The country of a Box is the applicant's State (that is, country) of residence if no State of	ity, full official designation. he address indicated in this dence is indicated below.) This person is: applicant only applicant and inventor inventor only (If this check-box is marked, do not fill in below.)			
State (that is, country) of nationality: GB	State (that is, country) of residence: GB			
This person is applicant all designated all designated				
Name and address: (Family name followed by given name; for a legal en. The address must include postal code and name of country. The country of Box is the applicant's State (that is, country) of residence if no State of resi	tity, full official designation. the address indicated in this dence is indicated below.) This person is: applicant only applicant and inventor inventor only (If this check-box is marked, do not fill in below.)			
State (that is, country) of nationality:	State (that is, country) of residence:			
1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	States except the United States the States indicated in the Supplemental Box			
Name and address: (Family name followed by given name; for a legal entity, full official designation. The address must include postal code and name of country. The country of the address indicated in this Box is the applicant's State (that is, country) of residence if no State of residence is indicated below.) This person is: applicant only applicant and inventor inventor only (If this check-box is marked, do not fill in below.)				
State (that is, country) of nationality:	State (that is, country) of residence:			
This person is applicant all designated for the purposes of:	d States except atte United States the States indicated in the Supplemental Box			
Further applicants and/or (further) inventors are indicated on another continuation sheet.				

Box No.V DESIGNATION OF STATES							
The following designations are hereby made under Rule 4.9(a)(mark the applicable check-boxes; at least one must be marked):							
Regional Patent							
X		P ARIPO Patent: GHGhana, GM Gambia, KE Kenya, LS Lesotho, MW Malawi, SD Sudan, SZ Swaziland, UG Uganda, ZW Zimbabwe, and any other State which is a Contracting State of the Harare Protocol and of the PCT					
X	EA	Eurasian Patent: AM Armenia, AZ Azerbaijan, BY Belarus, KG Kyrgyzstan, KZ Kazakhstan, MD Republic of Moldova, RU Russian Federation, TJ Tajikistan, TM Turkmenistan, and any other State which is a Contracting State of the Eurasian Patent Convention and of the PCT					
X	EP	European Patent: AT Austria, BE Belgium, CH and LI Switzerland and Liechtenstein, CYCyprus, DE Germany, DK Denmark, ES Spain, FI Finland, FR France, GB United Kingdom, GR Greece, IE Ireland, IT Italy, LU Luxembourg, MC Monaco, NL Netherlands, PT Portugal, SE Sweden, and any other State which is a Contracting State of the European Patent Convention and of the PCT					
X	OA	OAPI Patent: BFBurkina Faso, BJ Benin, CF Central African Republic, CG Congo, CI Côte d'Ivoire, CM Cameroon, GA Gabon, GN Guinea, GW Guinea-Bissau, MLMali, MR Mauritania, NE Niger, SN Senegal, TD Chad, TG Togo, and any other State which is a member State of OAPI and a Contracting State of the PC Til other kind of protection or treatment desired, specify on dotted line)					
BT-41	National Patent (if other kind of protection or treatment desired, specify on dotted line):						
×		Albania	K		Lesotho		
X	AM	Armenia	X	LT	Lithuania		
X	AT	Austria	X	LU	Luxembourg		
[32]	ΑIJ	Australia	[X]	LV	Latvia		
		Azerbaijan	X	MD	Republic of Moldova		
		2	=		Madagascar		
X		Bosnia and Herzegovina	X				
X		Barbados	X	MK	The former Yugoslav Republic of Macedonia		
X	BG	Bulgaria					
X	BR	Brazil	\boxtimes	MN	Mongolia		
X		Belarus	X	MW	Malawi		
		Canada	X	MX	Mexico		
⊠			X		Norway		
EX		and LI Switzerland and Liechtenstein	=		•		
X		China	X		New Zealand		
X	CU	Cuba	X	PL	Poland		
X	CZ	Czech Republic	\mathbf{X}	PT	Portugal		
X		Germany	X	RO	Romania		
X		Denmark	X	RU	Russian Federation		
[X]		Estonia	図	SD	Sudan		
_		Spain	Ø		Sweden		
X	ES		_				
X	FI	Finland	K		Singapore		
X	GB	United Kingdom	X	SI	Slovenia		
X	GD	Grenada	X	SK	Slovakia		
X	GE	Georgia	X	SL	Sierra Leone		
X	GH	Ghana	X	TJ	Tajikistan		
×		Gambia	X	TM	Turkmenistan		
X		Croatia	X		Turkey		
=		Hungary	×		Trinidad and Tobago		
ω Σ			K]		Ukraine		
X	ID	Indonesia	=				
X	IL	Israel	X		Uganda		
X	IN	India	X	US	United States of America		
X	IS	Iceland					
EX	JР	Japan	X	UZ	Uzbekistan		
X	KE	Kenya	X	VN	Viet Nam		
X		Kyrgyzstan	X	YU	Yugoslavia		
_		Democratic People's Republic of Korea	X		Zimbabwe		
X	I/L						
X	Check-boxes reserved for designating States (for the purposes of a national paths) which have become party to the PCT after the purpose of th						
		issuance of this sheet:					
X		Kazakhstan		DM	DOMINICA 🙎 ZA SOUTH AFRICA		
N N		LIC HAITED ADAD EMIDATES ALTZ TANZANIA					
X		CD COSTA DICA FT MA MODOCCO					
×	LR	Liberia	X	ĢĦ.	COSTA RICA . MA . MOROCCO		

Precautionary Designation Statement: In addition to the designations made above the applicant also makes under Rule 4.9(b) all other designations which would be permitted under the PCT except any designation(s) indicated in the Supplemental Box as being excluded from the scope of this statement. The applicant declares that those additional designations are subject to confirmation and that any designation which is not confirmed before the expiration of 15 months from the priority date is to be regarded as withdrawing the applicant at the expiration of that time limit. (Confirmation of a designation consists of the filing of a notice specifying that designation and the payment of the designation and confirmation fees. Confirmation must reach the receiving Office within the 15-month time limit.)

Sheet No. 4

Box No. VI PRIORITY C	LAIM	Further prio	rity claims are indicated	d in the Supplemental Box.
Filing date	Number		Where earlier applicat	ion is:
of earlier application (day/month/year)	of earlier application	national application: country	regional application:* regional Office	international application: receiving Office
item (1)		Country	Togrotian Ottico	receiving Office
11/02/1999 (11Feb1999)	9902970.4	GB		
item (2)				
item (3)				
of the earlier application(quested to prepare and tran s) (only if the earlier appli ternational application is t	ication was tiled with the the receiving Office)identif	ied above as item(s) (1)
* Where the earlier application is Convention for the Protection of I	an ARIPO application, it is n ndustrial Property for which i	mandatory to indicate in the S that earlier application was fi	Supplemental Box at least of led (Rule 4.10(b)(ii)). See	one country party to the Paris Supplemental Box.
Box No. VII INTERNATIO	ONAL SEARCHING AU	THORITY		
Choice of International Searce (if two or more International Sea competent to carry out the interna- the Authority chosen; the two-lette	arching Authorities are sear	quest to use results of earch has been carried out by one to the day/month/year)	rlier search; reference or requested from the Inter Number	e to that search (if an earlier rnational Searching Authority): Country (or regional Office)
ISA /				
Box No. VIII CHECK LIST	Γ; LANGUAGE OF FIL	ING		
This international applicatione the following number of sheet	te	nal application isaccompa	nied by the item(s) mark	ted below:
request : 4	1. X fee calcu	signed power of attorney		
description (excluding sequence listing part) : 45		general power of attorney;	reference number, if a	ny:
claims : 4		nt explaining lack of signat		
abstract : 1	_	document(s) identified in I		
drawings : 4		on of international applicat		
sequence listing part	7. 🔲 separate	indications concerning de	posited microorganism	or other biological material
of description : 2	8. X nucleotic	de and/or amino acid seque	ence listing in computer	readable form
Total number of sheets: 60	9. ☐ other (sp	pecify):		
Figure of the drawings which should accompany the abstract	•	anguage of filing of the ternational application:	English	
Box No. IX SIGNATURE	OF APPLICANT OR A	GENT		
Prouin &	name of the person signing and	the capacity in which the person	n signs (il such capacity is n	ot oovious from reading the reques
BILL, Kevi	IT			
1 Day 6 2 2 1 2 6 4		receiving Office use only		2. Drawings:
Date of actual receipt of the international application:				
Corrected date of actual re timely received papers or the purported international	drawings completing			received:
Date of timely receipt of the corrections under PCT Art	ticle [1(2):			not received:
5. International Searching Au (if two or more are compet	uthority ISA /		tal of search copy delay ch fee is paid.	red
Date of receipt of the record by the International Bureau:		emational Bureau use only	/	



PCT

INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference PHM 70478/W0	FOR FURTHER see Notification of (Form PCT/ISA/2	of Transmittal of International Search Report 220) as well as, where applicable, item 5 below.
International application No.	International filing date (day/month/year)	(Earliest) Priority Date (day/month/year)
PCT/GB 00/00357	08/02/2000	11/02/1999
Applicant ASTRAZENECA UK LIMITED		
according to Article 18. A copy is being to This international Search Report consists	_	
d. Designed the remove		
Basis of the report a. With regard to the language, the language in which it was filed, ur	International search was carried out on the baless otherwise indicated under this item.	asis of the international application in the
Authortty (Rule 23.1(b)).	was carried out on the basis of a translation of	
b. With regard to any nucleotide at was carried out on the basis of the	nd/or amino acid sequence disclosed in the l	nternational application, the international search
	onal application in written form.	
	ernational application in computer readable fo	rm.
_	o this Authority in written form.	
	o this Authority in computer readble form.	
International application	ibsequently furnished written sequence listing as filed has been furnished.	
the statement that the in furnished	formation recorded in computer readable form	is Identical to the written sequence listing has been
2. Certain claims were fo	und unsearchable (See Box I).	
3. Unity of Invention is la	cking (see Box II).	
4. With regard to the title,		
	submitted by the applicant.	
	shed by this Authority to read as follows:	
OLIGONUCLEOTIDE ARRAY	AND METHODS OF USE	
5. With regard to the abstract,		
X the text is approved as	submitted by the applicant.	
the text has been estable within one month from the	ished, according to Rule 38.2(b), by this Authone date of mailing of this international search r	ority as it appears in Box III. The applicant may, eport, submit comments to this Authority.
6. The figure of the drawings to be pu	blished with the abstract is Figure No.	
as suggested by the ap		None of the figures.
	alled to suggest a figure.	
because this figure bette	er characterizes the invention.	

wnational Application No PCT/GB 00/00357

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 C12Q1/68

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols) IPC 7 C12Q

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	WO 97 31256 A (BLOK HERMAN ;BARANY GEORGE (US); KEMPE MARIA (US); ZIRVI MONIB (US) 28 August 1997 (1997-08-28) see whole doc. esp. claim 13	1-5
X	EP 0 799 897 A (AFFYMETRIX INC) 8 October 1997 (1997-10-08) see whole doc., esp. claims 10,11	1-5
Υ	GB 2 312 747 A (ZENECA LTD) 5 November 1997 (1997-11-05) the whole document	6-16
Y	EP 0 416 817 A (ICI PLC) 13 March 1991 (1991-03-13) cited in the application see whole doc. esp. claims	6-16

Further documents are listed in the continuation of box C.	Patent family members are listed in annex.
"Special categories of cited documents: "A" document defining the general state of the art which is not considered to be of particular relevance "E" earlier document but published on or after the international filling date "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filling date but later than the priority date claimed	"T" later document published after the International filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art. "&" document member of the same patent family
Date of the actual completion of the international search 31 May 2000	Date of mailing of the international search report 07/06/2000
Name and mailing address of the ISA European Patent Office, P.B. 5818 Patentiaan 2 NL – 2280 HV Rijswijk Tel. (+31–70) 340–2040, Tx. 31 651 epo ni, Fax: (+31–70) 340–3016	Authorized officer Müller, F

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ernational Application No

		PCT/GB 00	7 00357
C.(Continu	ation) DOCUMENTS CONSIDERED TO BE RELEVANT		
Category °	Citation of document, with indication, where appropriate, of the relevant passages		Relevant to claim No.
X	WO 93 25563 A (HOPE CITY ; WALLACE ROBERT BRUCE (US)) 23 December 1993 (1993-12-23) see whole doc. esp. claims and figures		6-16
A	OLD J M: "DETECTION OF MUTATIONS BY THE AMPLIFICATION REFRACTORY MUTATION SYSTEM (ARMS)" METHODS IN MOLECULAR BIOLOGY, US, HUMANA PRESS INC., CLIFTON, NJ, vol. 9, 1991, pages 77-84, XP000775640 the whole document		·

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ternational Application No PCT/GB 00/00357

Patent documen cited in search rep		Publication date		Patent family member(s)	Publication date
WO 9731256	Α	28-08-1997	AU	2799797 A	10-09-1997
			CA	2244891 A	28-08-1997
			EP	0920440 A	09-06-1999
EP 0799897	Α	08-10-1997	NONE		
GB 2312747	A	05-11-1997	AU	2704997 A	26-11-1997
			EP	0896630 A	17-02-1999
			WO	9742345 A	13-11-1997
EP 0416817	A	13-03-1991	AT	144290 T	15-11-1996
			AU	622349 B	02-04-1992
			AU	6118490 A	14-03-1991
			DE	69028891 D	21-11-1996
			DE	69028891 T	13-03-1997
			GB	2235689 A,B	13-03-1991
			JP	3272686 A	04-12-1991
			NZ	235015 A	26-03-1992
			US	5525494 A	11-06-1996
WO 9325563	Α	23-12-1993	CA	2115342 A	23-12-1993
			AU	674127 B	12-12-1996
			AU	2251192 A	04-01-1994
			EP	0607151 A	27-07-1994
			JP	6509946 T	10-11-1994
			US	5981176 A	09-11-1999

Inter mai Application No PCT/GB 00/00357

A CLASSIFICATION OF SUBJECT MATTER IPC 7 C12Q1/68

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols) IPC 7 C12Q

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
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Date of the actual completion of the international search	Date of mailing of the international search report
31 May 2000	07/06/2000
Name and mailing address of the ISA European Patent Office, P.B. 5818 Patentiaan 2	Authorized officer
NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fex: (+31-70) 340-3016	Müller, F



Interr nal Application No PCT/GB 00/00357

		PCT/GB 0	0/00357
	ation) DOCUMENTS CONSIDERED TO BE RELEVANT		
Category *	Citation of document, with indication, where appropriate, of the relevant passages		Relevant to claim No.
(WO 93 25563 A (HOPE CITY ; WALLACE ROBERT BRUCE (US)) 23 December 1993 (1993-12-23) see whole doc. esp. claims and figures		6-16
	OLD J M: "DETECTION OF MUTATIONS BY THE AMPLIFICATION REFRACTORY MUTATION SYSTEM (ARMS)" METHODS IN MOLECULAR BIOLOGY, US, HUMANA PRESS INC., CLIFTON, NJ, vol. 9, 1991, pages 77-84, XP000775640 the whole document		
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inter		Application No
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Patent document cited in search report		Publication date	Patent family member(s)		Publication date
WO 9731256	A	28-08-1997	AU	2799797 A	10-09-1997
			CA	2244891 A	28-08-1997
			EP	0920440 A	09-06-1999
EP 0799897	Α	08-10-1997	NON		
GB 2312747	Α	05-11-1997	AU	2704997 A	26-11-1997
			EP	0896630 A	17-02-1999
			WO	9742345 A	13-11-1997
EP 0416817	Α	13-03-1991	AT	144290 T	15-11-1996
			AU	622349 B	02-04-1992
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			GB	2235689 A,B	13-03-1991
			JP	3272686 A	04-12-1991
_			NZ	235015 A	26-03-1992
			US	5525494 A	11-06-1996
NO 9325563	A	23-12-1993	CA	2115342 A	23-12-1993
			AU	674127 B	12-12-1996
			AU	2251192 A	04-01-1994
			EP	0607151 A	27-07-1994
			JP	6509946 T	10-11-1994
			US	5981176 A	09-11-1999